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Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

Applicant(s): Ganapathy et al. Serial No.: Unassigned

Express Mail No.: EV 201890140 US

Filed: Herewith

Docket: 275.00080101 Sheet 1 of 44

Figure 1

1/44

drIndyl sequence (tot_1 2602)

(ORF: 258 - 1976)

TTCACCGTTTCCGAATCGGACGAACCGGGCGTGATTGCTCTCCTGCTGCTTTCGAGATCGGAGTCCCGATAAGGATA TAACTACAACCTAAAGAGGAATCCAAGCCTCCTCCTGCCGCTAGTTTCGAAAAGTAAATAGAGTACTTGTTATCAAC TGGGAAGCGGAGATACATAGCTCCGATATTCCTGTGAAAGCCAGACAAACGGATACCAACGAACAATCGCCATATCT ACACGCCACCGCCACTGGACATCAAAATGGAAATTGAAATTGGCGAACAACCCCAGCCTCCGGTGAAGTGCTCCAAC TTCTTCGCTAACCACTGGAAGGGATTGGTTGTGTTCCTGGTGCCGCTGCTATGTCTGCCTGTTATGCTGCTAAACGA AGGCGCCGAATTTCGGTGCATGTACCTCCTTTTGGTAATGGCCATATTTTGGGTTACGGAAGCCTTGCCTCTATG TGACGTCCATGATACCGATTGTGGCCTTCCCAATAATGGGTATAATGAGCTCGGATCAGACTTGCCGCTTGTACTTC AAGGATACGCTGGTGATGTTCATGGGCGGCATTATGGTCGCCCTGGCTGTGGAGTACTGTAATCTACACAAACGTCT TGCCTTGAGGGTAATCCAGATCGTGGGCTGCAGTCCCCGCAGATTACACTTTGGCCTCATCATGGTTACAATGTTTT TGAGCATGTGGATTTCGAACGCCGCCTGTACTGCCATGATGTGTCCGATTATCCAAGCCGTGCTGGAGGAGCTGCAG GCTCAGGGTGTCTGCAAAATCAACCATGAGCCTCAATACCAAATCGTTGGAGGCAACAAGAAAAACAACGAGGATGA GCCACCATACCCCACCAAGATCACTCTGTGCTACTATCTGGGCATTGCCTACGCCTCCTCGCTGGGTGGCTGTGGAA CCATCATCGGAACTGCCACCAATCTTACCTTCAAGGGCATCTACGAGGCTCGTTTCAAGAACTCCACCGAACAGATG GACTICCCCACCTICATGTTCTACTCGGTGCCATCCATGTTGGTCTACACCTTGCTGACATTCGTGTTCCTGCAATG GCACTTCATGGGTCTGTGGCGTCCCAAGAGCAAGGAGGCACAGGAAGTCCAGAGGGGGACGAGAGGGCGCCGATGTCG CCAAAAAGGTTATCGATCAGCGCTACAAGGATCTGGGTCCCATGTCCATTCACGAGATCCAAGTGATGATTCTGTTC ATTTTTATGGTTGTGATGTACTTCACCCGCAAGCCCGGCATCTTTTTGGGATGGGCCGATTTGCTGAATTCCAAGGA CATTCGTAACTCTATGCCCACTATTTTTGTCGTCGTCATGTGCTTCATGCTGCCCGCCAATTATGCTTTCCTACGCT ACTGCACCAGACGCGGTGGTCCAGTGCCCACGGGTCCCACTCCATCGCTGATCACCTGGAAGTTCATCCAGACCAAG GTGCCATGGGGTCTGGTGTTCCTGCTTGGCGGTGGCTTCGCTTTGGCCGAAGGCAGCAGCAGCAGCGGCATGGCCAA GCTGATTGGCAATGCTCTGATTGGATTGAAGGTTCTGCCCAACTCTGTCCTCTTACTGGTGGTCATCCTGGTGGCTG TGTTCCTGACCGCCTTCAGCTCCAATGTGGCGATTGCCCAACATTATTATTCCCGTTCTGGCCGAGATGTCCCTGGCC ATTGAGATCCATCCTGTACCTGATCCTGCCCGCTGGCTTGGCCTGCAGTATGGCCTTCCACCTGCCGGTTAGTAC TCCGCCCAACGCTTTGGTTGCTGGCTATGCCAACATTAGGACGAAGGACATGGCCATTGCTGGAATCGGTCCGACCA TCATTACCATCATCACCCTGTTTGTTTTCTGCCAAACCTGGGGCCTGGTCGTCTATCCGAACCTTAACTCGTTCCCC **AACATACCCGTCACAGCGATAAAGTTGAGGAAAATTTAGGGAATTTTAAACGAAAAGTGCCTTTGCTGACAGCGAAA** AATGTGAAAAATATTTAACTATGTATACTTGCATTTCAGAGTTGCGAAAAGTTTTGATACAAAAGCATTACCTACTG TTTAGAAAAATGTGTTAAAAAAAAAACGTATCGCAATATACTGTTAATCAGGAATTGAACACCTGGTCTACGCACTC AGCTAAATATTTAAATACAAATTAATGTTACTTAATTGTTGCATTTAGCATAAAAATGGAAAAGATTTGGAAAAGTT AGAACAGTTTGTTCAATGGCAGCCCTGGCCTGATATTTTTAAATAACTAGACTGAGAGAACTTACATATTCATAC TAGCGGTAGGCTAAGCTTAAATGATACTGTGTACATTTTCAGATGATTTATGTTTTATATAGTTTGTAAAAAAATATT

SEQ ID No:1

Peptide seq (total 572)

MEIEIGEQPQPPVKCSNFFANHWKGLVVFLVPLLCLPVMLLNEGAEFRCMYLLLVMAIFWVTEALPLYVT SMIPIVAFPIMGIMSSDQTCRLYFKDTLVMFMGGIMVALAVEYCNLHKRLALRVIQIVGCSPRRLHFGLI MVTMFLSMWISNAACTAMMCPIIQAVLEELQAQGVCKINHEPQYQIVGGNKKNNEDEPPYPTKITLCYYL GIAYASSLGGCGTIIGTATNLTFKGIYEARFKNSTEQMDFPTFMFYSVPSMLVYTLLTFVFLQWHFMGLW RPKSKEAQEVQRGREGADVAKKVIDQRYKDLGPMSIHEIQVMILFIFMVVMYFTRKPGIFLGWADLLNSK DIRNSMPTIFVVVMCFMLPANYAFLRYCTRRGGPVPTGPTPSLITWKFIQTKVPWGLVFLLGGGFALAEG SKQSGMAKLIGNALIGLKVLPNSVLLLVVILVAVFLTAFSSNVAIANIIIPVLAEMSLAIEIHPLYLILP AGLACSMAFHLPVSTPPNALVAGYANIRTKDMAIAGIGPTIITIITLFVFCQTWGLVVYPNLNSFPEWAQIYAAAALGNKTH

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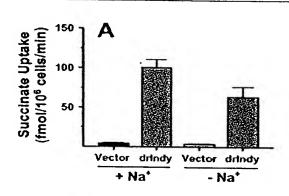
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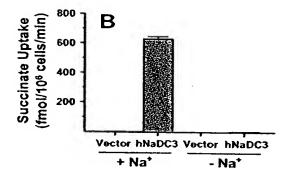
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Docket: 275.00080101 Sheet 2 of 44

Figure 2



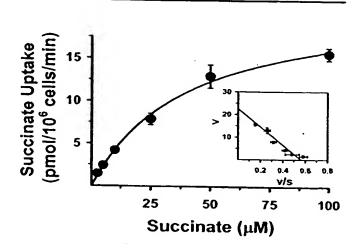


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Docket: 275.00080101 Sheet 3 of 44

Figure 3

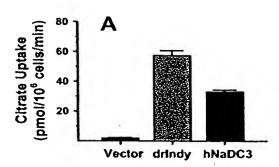


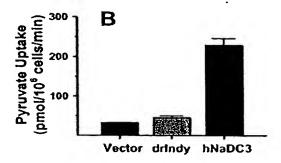
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Serial No.: Unassigned Filed: Herewith Express Mail No.: EV 201890140 US

Docket: 275.00080101 Sheet 4 of 44

Figure 4





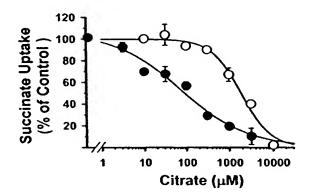
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Docket: 275.00080101 Sheet 5 of 44

Figure 5



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Serial No.: Unassigned Filed: Herewith Express Mail No.: EV 201890140 US

Docket: 275.00080101 Sheet 6 of 44



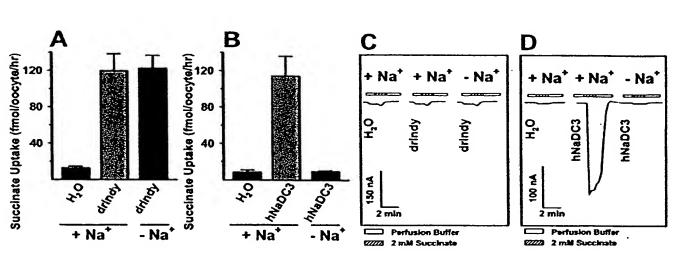


Figure 6

Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

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Docket: 275.00080101 Sheet 7 of 44

7/4

Figure 7

Rat INDY seq. (3191 nt + 63 nt polyA)

CCAGTCTGTCTCCCTTTCACGCGATGGCTTCGGCGAAGACTTATGTGACCAAGTTCAAGTCCTTCGTGATTTTGTTCTTCGCCCC GATCCTGCTGCTTCCACTCATCATTCTGGTACCTGACAAGTTTGCCAGGTGTGCCTATGTTATAATCCTCATGGCCATCTACTGG TATGTGTCCAATACATGACGGACACCAACATGCTGTTCCTGGGCAGTCTCATTGTGGCCACGGCTGTGGAACGTTGGGAAC7TCA TAAGAGGATTGCTCTGAGAATGCTACTCTTTGTGGGGACCAAGCCTTCACGGCTGATGCTGGGCTTCATGTTCGTCACAGCCTTC CTGTCCATGTGGATCAGCAATACTGCCACCACGGCCATGATGATACCCATTGTGGAGGCCATGCTGGAGCAAATGGTAGCCAACA TGAAGACCCCAGTGTGCAGAAGCAGGAGGATGAAGAAACAAAGAATATGTACAAGGCTATGAACCTATGTGTGCTACGCAGCC **AGCATCGGGGGTACAGCACCTTGACCGGGACGGGACCCAACGTGCTGCTGCTGGGCCAGATGCAGGAATTGTTTCCTGACAGAA AAGACGTCATGAACTTTGCATCTTGGTTTGCCATTTGCCCTCCCAAACATGCTTTTGATGCTGGTGATGGCCTGGCTGTGGCTCCT** GTGTTTTTP.CATGAGACCCAATTTAAAAAAAACTTGCATCTGCTGGGGGAAGAAGAAGAAGGACACGGAGAAGATTCCCTAAG GTGCTGTATGAGGAGTACAGGAAGCTGGGGCCCTTGAGCTACGCTGAATGCAACGTGCTCTTTTGCTTCGGCCTGCTCATCATCC CACAGTGGCCATCTTTGTGGCCATTTTGCTTTTCATCGTACCCTCACAAAAGCCCCAAGTTCAATTTCAGCCGCCAGACTGAGGAA GAAAGGAAAACTCCCTTCTACCCCCCGCCACTGCTGAATTGGAAAGTCACCCAAGAGAAAGTGCCCTGGGGCATTGTGCTGCTC **AGTGAGACCTGCTATTATTACCTTGATCTTGTCCTGTATTGTTGCAATGACCACAGAGTGCACGAGTAACGTGGCCACTACTACC** CACTTGCCTTCATGTTGCCTGTGGCCACCCCACCTAACGCCATCGTGTTTGCCTACGGACACCCTCAAAGTTATTGACATGGTAAA ANATTCCCTGACTGGGCAAATTTGACACATATTAACACTTAGGAGAACCACAAGAGCACAGGCTTGTCCCCCAACCCTTTCGAGG ACTGCGAACCTTCTGGCACACCTTGCACAGAGCACTGGTGCTCATACCCCAGTGTGACCCCAATGATGTCAACACCCCAAGAAGATGT CCTTTGAGAGGTCGTGAGGCCCATCTTCCCTAGGACCCTTCCATCTCACCTGGGCAGAAACAGAGGGACTGGGGCTCAAGTCCT GTACCACGTGGCTTTGAAAGACTTCTGACTCCATGCTGGGCTCTGGTTCTCACATGCCTGTTCCCACGGTCTCCACATGGGGATC AGATGACCAGGAGACAGCCCCTGTGCCTCTTCTGGATGTTCCCAGATCACCATCTCTATCACCAAGGAAGAACTTCCTCTCCAGG ACAGAACTCTGATCTTGAACACTTTCCACTGCCAGAGTTAGAGTGGAAATCACGGCCCCTGAAGACTTTGACTCTACATGGTGC CATCTCCAACCACTGGGCAACCTGAGGTGCTAACATTGAGGCCTTCCTGCTCACCCTTGGCTGACCTGTTCCCTACTTGCCTTAT . CTCTTATTAGTTAACAGTTLGAGGCCCCTTCCCAGCTCCCAGLGAGACTTCATCAACTCCTAGATGCTCCTGGCTGAGGCTCCC CAGAAAGTCCCTGTTTGATTGTTCTGGGGATGCTCATCCTGCCAGCCTGGAGCAGCTGGGTCATACATCAGGGATGGACAATGT TRAGGGCTGGGCCACCACAGTTGGTCCTGTTGGACTCTTGAGCTCCTCCAGAAGGCCTTTTCCTTGCTGTGTGCACTGTGGGC GGT&G>GTCCATGGGGA&CCCATGGGACCACTCACATGAAAGGGAGAGAGAGAGAAGATCTCCCTTGTCCTTCAG<mark>GGATGC</mark>T CTCTTCCTTGCTTAATTTGCTCTGAAAAGAAGCATGAGTGGGGAGATAAGATCCGTGGATGTCATTTTATTTTTCCAGGCAGAAG AAGGCTTGCTCTTGTCAAATACTCTGTCTGCTATGGAAAGTTCCAGTGTGCTGACTGGTCTGTGATTTCGTGCCTTGTGAAGGAG **AAAAAAAAAAAAAAAAAAAA** SEQ ID NO:3

(572 aa)

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Serial No.: Unassigned Express Mail No.: EV 201890140 US Filed: Herewith

Docket: 275.00080101 Sheet 8 of 44

844



Figure 8

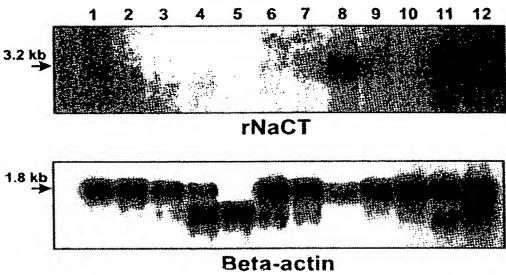
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Docket: 275.00080101 Sheet 9 of 44

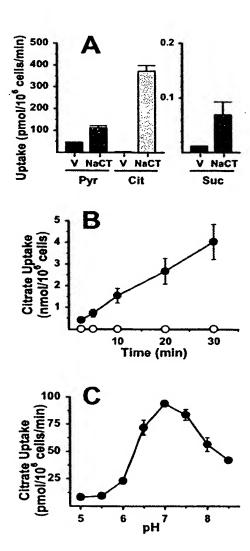
Figure 9



Docket: 275.00080101 Sheet 10 of 44

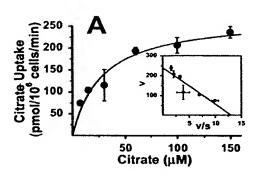
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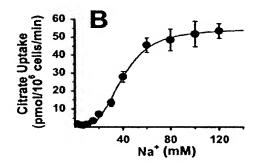
Figure 10

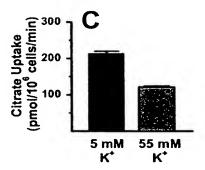


Docket: 275.00080101 Sheet 11 of 44

Figure 11

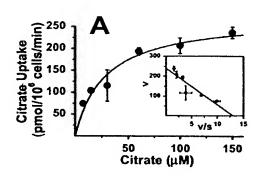


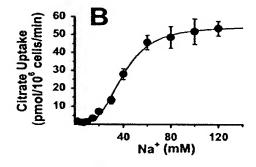


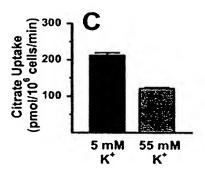


Docket: 275.00080101 Sheet 12 of 44

Figure 12

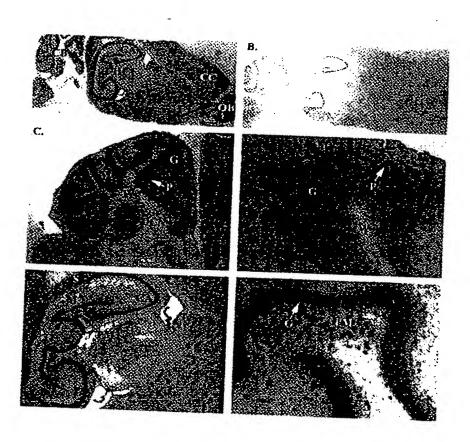






Docket: 275.00080101 Sheet 13 of 44

Figure 13



Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

Applicant(s): Ganapathy et al.

Serial No.: Unassigned Express Mail No.: EV 201890140 US Filed: Herewith

Docket: 275.00080101 Sheet 14 of 44

Figure 14

Human NaCT sequence

14/44

(3207 nt + 41 nt polyA) ORF: 13 -1719 (total 1707 nt)

CCCTCCCGCGCG

ATGGCC?CGGCGCTGAGCTATGTCTCCAAGTTCAAGTCCTTCGTCATCTTGTTCGTCACCCCGCTCCTGCTGCTGCCACTCC **CATTCTGATGCCCGCCAAGTTTGTCAGGTGTGCCTACGTCATCATCCTCATGGCCATTTACTGGTGCACAGAAGTCATCCC ATGAAGGACACCAACATGCTGTTCCTGGGCGCCTCATCGTGGCGCTGTGGAACCTGGAACCTGCACAAGAGAACATC CCCTGCGCACGCTCCTCTGGGTGGGGGCCAAGCCTGCACGGCTGATGCTGGGCTTCATGGGCGTCACAGCCCTCCTGTCCAT GTGGATCAGTAACACGGCAACCACGGCCATGATGGTGCCCCATCGTGGAGGCCATATTGCAGCAGATGGAAGCCACAAGCCGCA GCCACCGAGGCCGGCCTGGAGCTGGTGGACAAGGGCCAAGGCCAAGGAGCTCAAGTGATTTTTGAAGGCCCCCA CTCTGGGGCAGCAGGAAGACCAAGAGCGGAAGAGGTTGTGTAAGGCCATGACCCTGTGCATCTGCTACGCGCCAGCATCGG GGCACCGCCACCCTGACCGGGACGGGACCCAACGTGGTGCTCCTGGGCCAGATGAACGAGTTGTTTCCTGACAGCAAGGAC CTCGTGAACTTTGCTTCCTGGTTTGCCATTTGCCTTTCCCAACATGCTGGTGATGCTGCTGTTCGCCTGGCTGTGGCTCCAGT GCTGCAGGAGGAGTACCGGAAGCTGGGGGCCCTTGTCCTTCGCGGAGATCAACGTGCTGATCTGCTTCTTCCTGCTGGTCATC ATGCCACTGTGGCCATCTTTGTGGCTACCCTGCTATTCATTGTGCCTTCACAGAAGCCCAAGTTTAACTTCCGCAGCCAGAC GTGCTGCTACTAGGGGGGGGGATTTGCTCTGGGCTAAAGGATCCGAGGCCTCGGGGGCTGTCCGTGTGGATGGGGAAGCAGATGG AGCCCTTGCACGCAGTGCCCCCGGCAGCCATCACCTTGATCTTGTCCTTGCTCGTTGCCGTGTTCACTGAGTGCACAAGCAA CGTGGCCACCACCACCTTGTTCCTGCCCATCTTTGCCTCCATGTCTCGCTCCATCGCCTCAATCCGCTGTACATCATGCTG CCCTGTACCCTGAGTGCCTCCTTTGCCTTCATGTGCCTGTGCCACCCTCCAAATGCCATCGTGTTCACCTATGGGCACC TCAAGGTTGCTGACATGGTGAAAACAGGAGTCATAATGAACATAATTGGAGTCTTCTGTGTGTTTTTTGGCTGTCAACACCTG GGGACGGGCCATATTTGACTTGGATCATTTCCCTGACTGGGCTAATGTGACACATATTGAGACTTAG GAAGAGCCACAAGACCACACACACAGCCCTTACCCTCCAGGACTACCGAACCTTCTGGCACACCTTGTACAGAGTTTTGG TGCAGAGATGGACATGGGCAGCTGGAGGGTAGGCTCAGAAATGAAGGGAACCCCTCAGTGGGCTGCTGGACCCATCTTTCCC AAGCCTTGCCATTATCTCTGTGAGGGAGGCCAGGTAGCCGAGGGATCAGGATGCAGGCTGCTGTACCCGCTCTGCCTCAAGC ATCCCCCACACAGGGCTCTGGTTTTCACTCGCTTCGTCCTAGATAGTTTAAATGGGAATCAGATCCCCTGGTTGAGAGCTAA GACAACCACCTACCAGTGCCCATGTCCCTTCCAGCTCACCTTGAGCAGCCTCAGATCATCTCTGTCACTCTGGAAGGGACAC CCCAGCCAGGGACGGAATGCCTGGTCTTGAGCAACCTCCCACTGCTGGAGTGGGAATCAGAGCCTCCTGAAGCCTC AGTGCACCACGCCCAACCTACGCCCTTCATCACTTGGTTCTGTTTTAATCGACTGGCCCCCTGTCCCACCTCTCCAGTGAG TICTCCCAGGCAGGTCATCTTTTCTGGGAGATGCTTCCCCTGCCATCCCAAATAGCTAGGATCACACTCCAAGTATGGGCA ACTGGCTATGCCACTTCAGAGTCTTTCATGCCAGCGTTTGAGCTCCTCTGGGTAAAATCTTCCCTTTGTTGACTGGCCTTCA CAGCCATGGCTGGTGACAACAGAGGATCGTTGAGATTGAGCAGCGCTTGGTGATCTCTCAGCAAACAACCCCTGCCCGTGGG CCANTCTACTTGAAGTTACTCGGACAAAGACCCCAAAGTGGGGCAACAACTCCAGAGAGGCTGTGGGAATCTTCAGAAGCCC CCCTGTAAGAGACAGACATGAGAGACAAGCATCTTCTTTCCCCCCGCAAGTCCATTTTATTTCCTTCTTGTGCTGCTCTGGAA ALGGACCGAGTATGTGTGGGTTGGGTGGGACGATTCCTGACCACACTGTCCAGCTCTTGGTCTCATTAPATGCTCTG

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SEO ID No:2

Amino acid sequence (568 aa)

MASALSYVSKEKSEVILEVTPLLLLPLVILMPAKEVRCAYVIILHAIYHCTEVIPLAVTSLMPVLLEPLEQILDSRQVCVQY
MKDTNMLFLGGLIVAVAVERNMLHKRIALRTLLWVGAKPARLMLGFMGVTALLSMHISNTATTAMMVPIVEAILQOMEATSA
ATEAGLELVDKGKAKELPGSQVIFEGPTLGQQEDQERKRLCKAMTLCICYAASIGGTATLTGTGPNVVLLGQMBELFPDSKD
LVNFASHFAFAFPNMLVMLLFAWLHLQFVYMRFNFKKSHGCGLESKKNEKAALKVLQEEYRKLGPLSFAEINVLICFFLLVI
LXFSRDPGFMPGHLTVAWVEGETKYVSDATVAIFVATLLFIVPSQXPKFNFRSQTEERKTPFYPPPLLDHKVTQEKVPWGI
VLLLGGGFALAKGSEASGLSVHMGKQMEPLHAVPPAAITLILSLLVAVFTECTSNVATTTLFLPIFASMSRSIGLNPLYIML
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Filed: Herewith

Docket: 275.00080101 Sheet 15 of 44

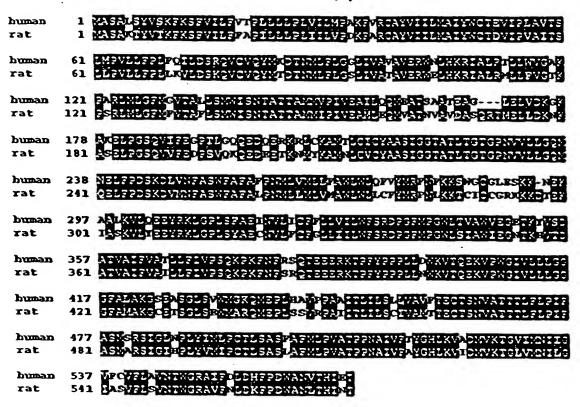
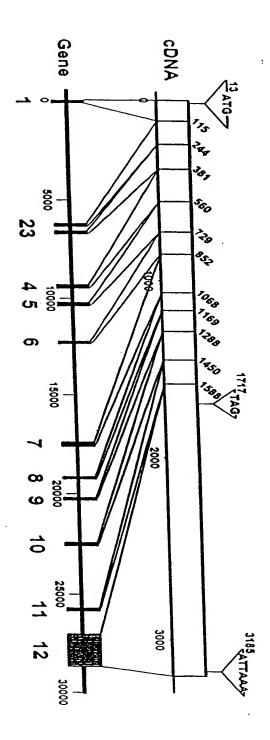


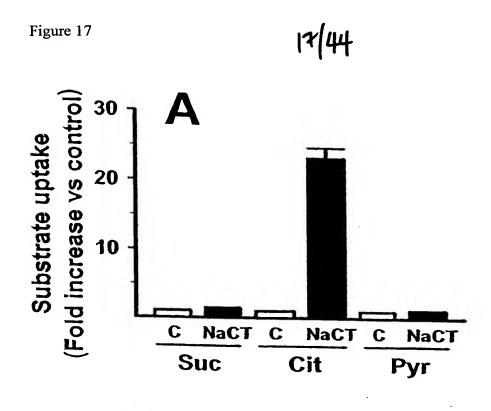
Figure 15

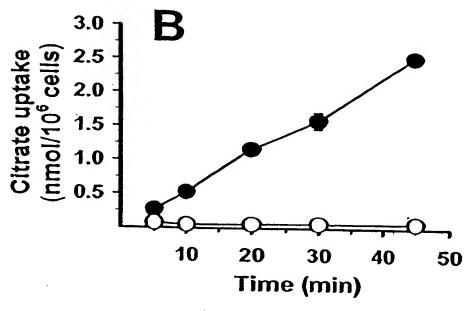
Docket: 275.00080101 Sheet 16 of 44

Figure 16

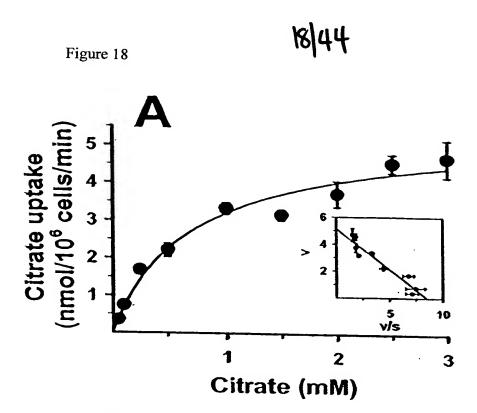


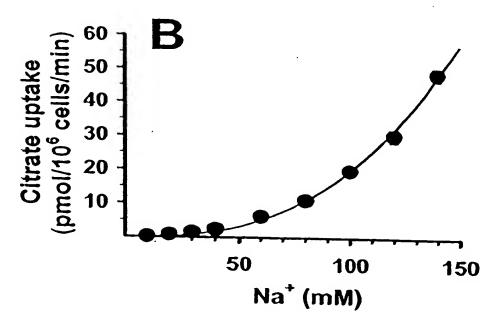
Docket: 275.00080101 Sheet 17 of 44





Docket: 275.00080101 Sheet 18 of 44





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Serial No.: Unassigned Filed: Herewith

Express Mail No.: EV 201890140 US

Docket: 275.00080101 Sheet 19 of 44



Figure 19

ceNaCT cDNA & Protein Sequences

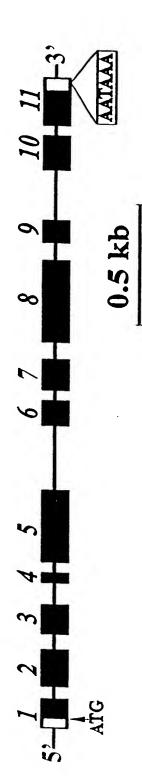
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•		100
101	M K P S P Q R T L I K K L L V L L G P L V A V P CTCTACTCTTTTTTGGACCGGAGTACAGATGTCTTTTCTCAATAATATTTTTATCAACCTATTGGAGAAGCATTTCCCATTGGTGTCACTTCTCT	
201	CITICCATTOGCACTITATCCAATTCTTCAGATTGTTCCATCTCAAGCAAATTAGTCAGTTTTTTAAACTAATTTGTTCATTTTTTTT	200
201		300
301	FPLALYPILQIVPSKQISPVYPKDSIVLPMCTL AGCATGGCGGTAGAAGCAACTGGACTCCATCGAAGAATCGCACTGAAATTATTAACAAAAGTTGGAGCAAAGCAACCAGTAATGCTGCTGGGT	
401	S M A M A V E A T G L H R R I A L R L L T R V G A R Q P V M L L G P TCATGTGCATCACGAGTTTCATATCATTTTTCGTTTCTGACACAGCATGCACAGCTCTTATGTGTCCAACCGCTGTGGCACTCCTGATGAGTATGTCTGA	400
501	TOCAGTTCAACATTGAAGAAGACAGCAGGAAGCCAACAACATCATCTACTTCATCTTCTCAACATTGAACATCATCAGGAAGCCAACAACATCATCTACTTCTTCTTCTTCTTCTTCTTCTT	500
501	A V Q H L K E D H R K P K P P P D D A T V A E K H R I D D M T P Q	600
601		700
701	GAGAAAATATTCACAAGAGATACCCCGAGGGACAAGTGACAATGCATACCTACAATGGTTTTCGCGATACCACCAATGTTTGTCTACCTTCTTGC	
801	ATCTITATATIATICTOGTOGTIATIATICGCCCCTCGCACTITICGCTCTCTCTAGACCCCCTCCCACTTTCAAACCCCCCCCC	800
	S Y I I L V C Y F M G P S T F A R W F E R P S K E E A H L K K L I	900
901		1000
1001	E K N I Q T M Y E D L G D V S W G E K S V F V F P I L L I G S W I S CTCGTGATCCCGGATTCACACCCGGATCTTTTACCACACAGAAACTTCATATCAGACAGTGTTTCTGGAGTCTTGATTTCTTATTTAT	
1101	R D P G P T P G W G D L L P H R N F I S D S V S G V L I S C I L F TOTTTOGCCCAMAGATCATTTGATCAMTGATCATTGATTCAMTGATCATTAMTGATTAMTGATCATTAMTGATCATTAMTGATCATTAMTGATCATTAMTGATCATTAMTGATCATTAMT	1100
	V W P K D P F D P I D P H A P I L K W T D H K S K F S W S C T L L ATCOGTGCTGGGTATGCTATTTCAGAAGGAGTTGATAAATCGGGATTATCTAGATTGATT	1200
1201	I G A G Y A I S E G V D K S G L S B L I S C C V T V	1300
1301	TOTALITATION TOTALI	
1401	TGCTGANTCANTGGGGTTCATCTTTATCTTGCTCTTCCACTACTGTGGTTTSTTCATTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTTGTT	1400
	A E S M G V M P L Y L A L P T T V A C S F A F M L P I S T P P N A	1500
1501	V V Y D T K V I S M V E M I V C C P I I N I S C T I N I S C T I N I S	1600
1601	TICCCCAAAATTTTGTAATATCTTCAGAAAACTCTTCATATCCAGTTTGCTAATTTTTGTACAAAAT	1700
1701	W T Y P I F S L N I P P E N I V I S S E N S S Y P V C . GTGTATTGTCCGAATGAAACGTGTATTTATTAAAAAAAAA	

Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION Applicant(s): Ganapathy et al.

Serial No.: Unassigned Filed: Herewith

Express Mail No.: EV 201890140 US

Docket: 275.00080101 Sheet 20 of 44



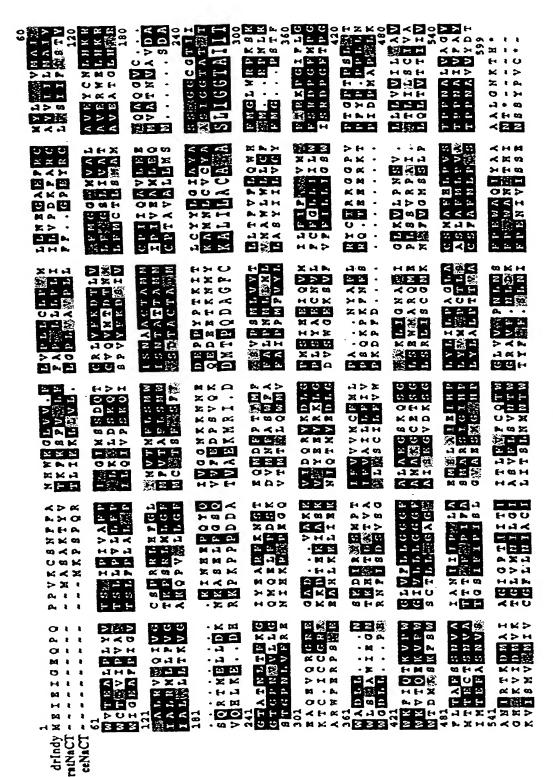
Serial No.: Unassigned

Express Mail No.: EV 201890140 US

Filed: Herewith

Docket: 275.00080101 Sheet 21 of 44

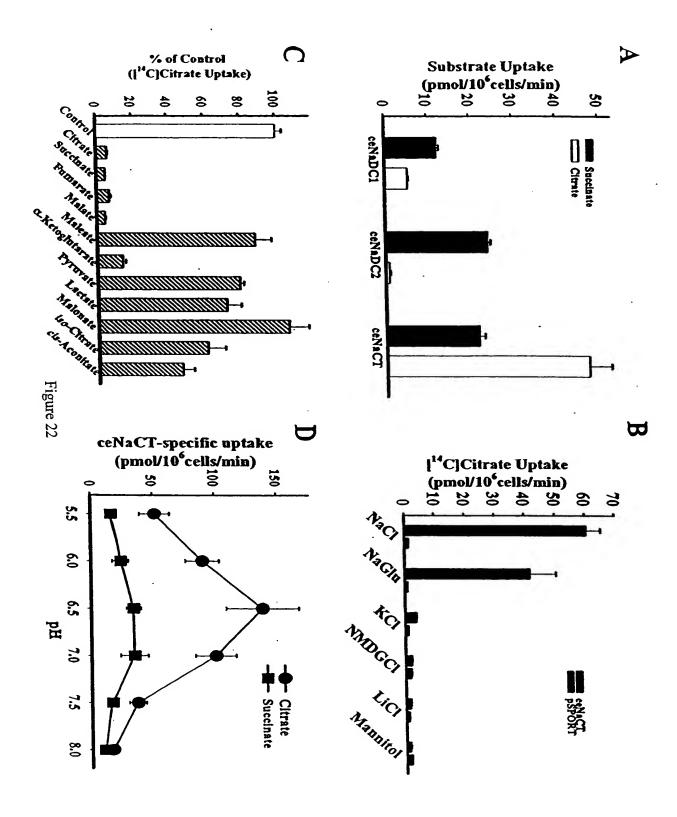
21/44



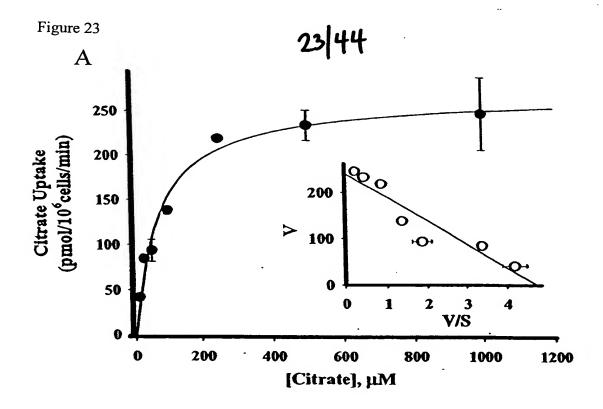
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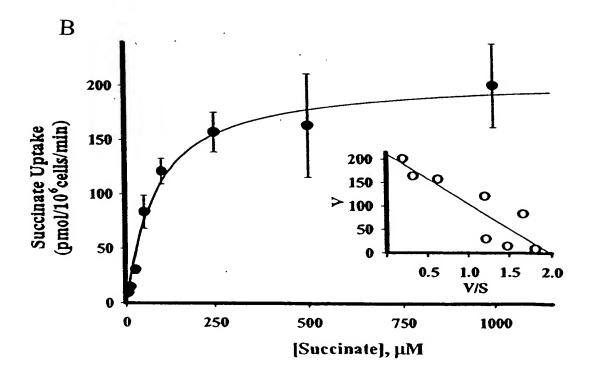
Figure 7

Docket: 275.00080101 Sheet 22 of 44

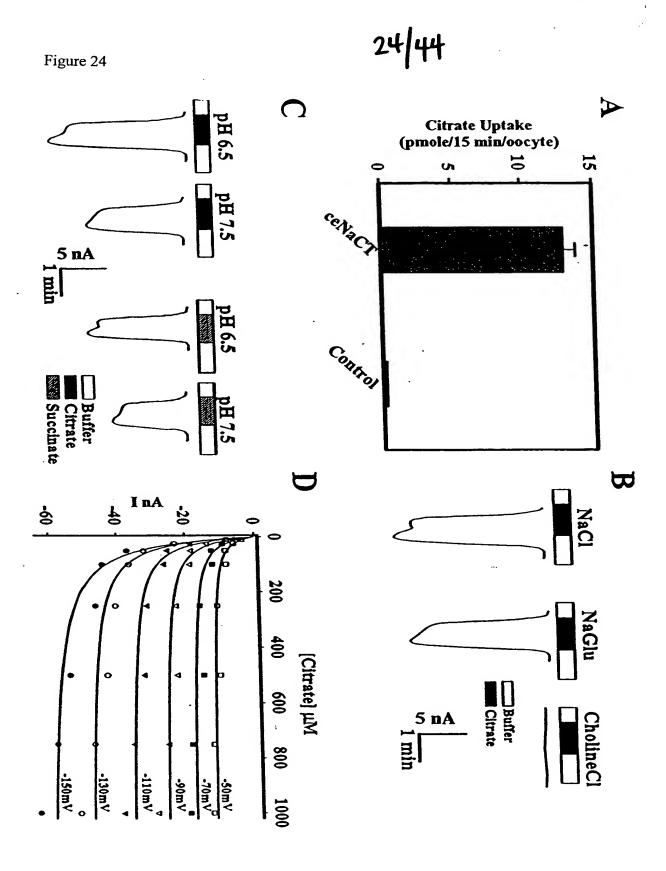


Docket: 275.00080101 Sheet 23 of 44





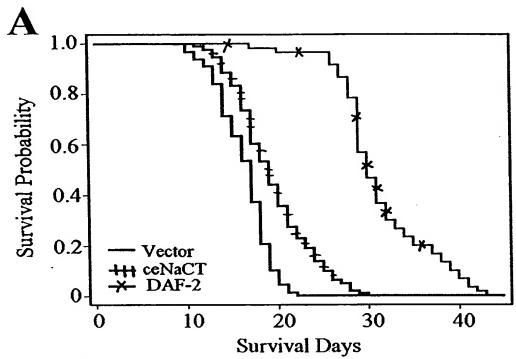
Docket: 275.00080101 Sheet 24 of 44

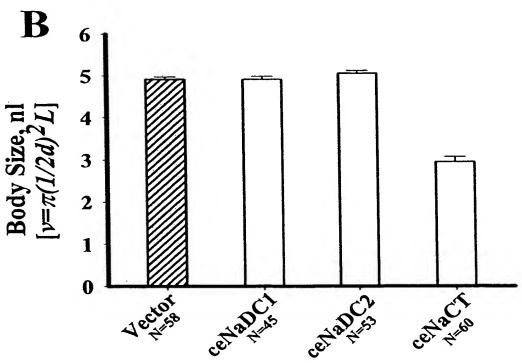


Docket: 275.00080101 Sheet 25 of 44









Docket: 275.00080101 Sheet 26 of 44

26/44

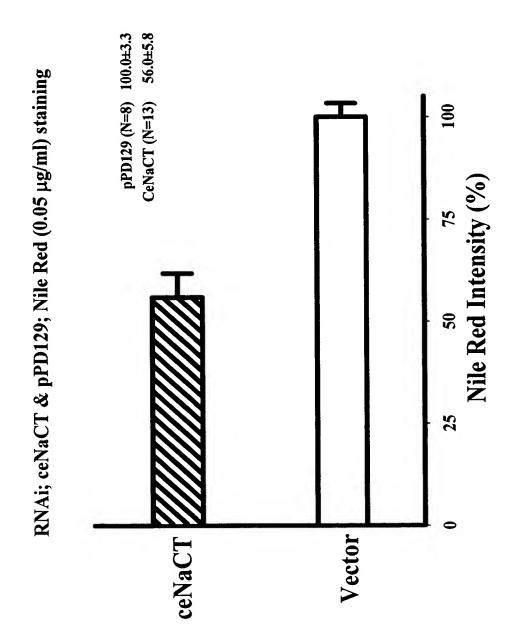


Figure 26

TIME: NACT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION Applicant(s): Ganapathy et al.

Serial No.: Unassigned
Express Mail No.: EV 201890140 US

Filed: Herewith

Docket: 275.00080101 Sheet 27 of 44

27/44

Figure 27

SEQ ID NO:9

Mouse NaCT sequence

cDNA sequence (16 nt + 1719 nt)

GTCTCCCTTTCACGCG

ATGGATTCGGCGAAGACTTGTGTGACCAAGTTCAAGTCCTTTGCGATTTTGCTCTTCACCCCGATCCT GATGCTTCCACTCGTCATTCTGATACCTGACAAGTTTGCCAGGTGTGCCTATGTTATAGTCATTATGG CTITTGAAGGTTCTGGACTCCAAGCAGGTATGTATCCAATACATGAAGGACACCAACATGCTGTTCCT GGGCAGTCTCATTGTGGCTGTGGGAACGCTGGAAACTTCATAAGAGGGTTGCCCTGAGAATGC-TGCTCTTTGTGGGGACCAAGCCCTCACGGCTGATGCTGGGCTTTATGTTTGTCACGGCCTTCCTGTCC ATGTGGATCAGCAATACTGCCGCCACAGCCATG ATGATACCCATTGTGGAGGCCATGCTGCAGCAAAT GATAGCCGCCAATACAGCTGTGGAGGCCCAGCCTGGGGCACACTGGAGCTGCTGGACAAGAACAAGACCA GCGAGTTGCCAGGAAGCCAGGTGGTATTTGAAGACCCCAATGTGCAGGAGCAGGAAGACGAAGAACA GACCGGGACGGACCCAACGTGGTGCTCCTGGGCCAGATGCAGGAATTGTTTCCTGACAGTAAAGATG TCCTGAACTATGCATCTTGGTTTGGATTTGCCTTCCCCAACATGGTGATGATGCTGGTGCTGGCCTGG GAGGGACACCGAGAAGATTGCCTACAAAGTGCTGAACGAGGAGTACCAGAAGCTGGGGTCCTTGAGCT ACCCTGAATGCAACGTGCTCTTTTGCTTCACCCTACTTGTCATCCTGTGGTTCTCCCGAGACCCCGGC TTCATGCCTGGCTGGCTGATTCGCCTGGGTCGAGGGAAACACCGTTCATATCACAGATGCCACAGT GGCCATCTTTGTGGCCATTTTGCTTTTCATCATACCTTCACAAAAGCCCCAAGTTCAACTTCAGCAGCC AGACTGAGGAAGAAAGGAAAACTCCGTTCTACCCCCCAGCACTGCTGGATTGG AAAGTCGCCCAAGAG AAAGTGCCCTGGGACATCGTGCTCCTGGGGGGGGGGGTTTGCTATGGCAAAAGGATGTGAGACGTC AGGGCTCTCGAAGTGGATGGCAGCAGATGGAACCCTTGAGATTAGTGAAACCTGCTGTCATTACCT TGATCTTGTCCTGTCTTGCCAATGACCACAGAGTGCACAAGTAACGTGGCCACTACCACCCTGTTC CTGCCTATCTTTGCCTCCATGGCTCGTTCCATTGGTA TCCATCCTCTGTATGTCATGATTCCCTGTAC CATGAGTGCTTCACTTGCCTTCATGTTGCCTGTGGCCACCCCACCGAATGCCATCGTGTTTGCCTACG GACACCTCAGAGTTGTTGACATGATGAAAACAGGATTGATAATGAACTTCGTTGGAATCCTATCTGTG TTTCTGTCAGTCAACACCTGGGGTCGGGCTATGTTTAACTTGGATAACTTCCCCGACTGGGCAAATTC **AACAÁGTGTTAACACTTAG**

Protein sequence (572 nt)

SEQ ID NO:10

MDSAKTCVTKFKSPAILLFTPILMLPLVILIPDKFARCAYVIVIMAVYWCTDVIPVAVTSLLPVLLFPLLKVLDSKQV
CIQYMKDTNMLFLGSLIVAVAVERWKLHKRVALRMLLFVGTKPSRLMLGFMFVTAPLSMWISNTAATAMMIPIVEAML
QQMIAANTAVEASLGTLELLDKNKTSBLPGSQVVFEDPNVQEQEDEBTKNMYKAMHLCVCYSASIGGTATEGTGPNV
VLLGQMQELFPDSKDVLNYASWFGFAFPNMVMMLVLAWLWLQCLYMRHNLKKTCICCGBKKRDTEKIAYKVLNBBYQK
LGSLSYPECNVLFCFTLLVILWFSRDPGFMPGWLSFAWVEGNTVHITDATVAIFVAILLFIIPSQKPKFNFSSQTEBB
RKTPFYPPALLDWKVAQEKVPWDIVLLLGGGFAMAKGCBTSGLSKWMAAQMEPLRLVKPAVITLILSCLVAMTTECTS
NVATTTLFLPIFASMARSIGIHPLYVMIPCTMSASLAFMLPVATPPNAIVFAYGHLRVVDMMKTGLIMNFVGILSVFL
SVNTWGRAMFNLDNFPDWANSTSVNT

Applicant(s): Ganapathy et al. Serial No.: Unassigned Express Mail No.: EV 201890140 US

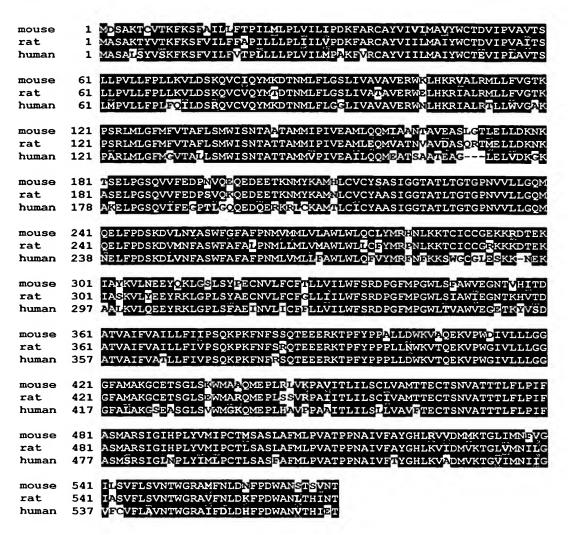
Filed: Herewith

Docket: 275.00080101 Sheet 28 of 44

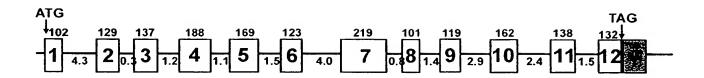
Figure 28

28/44

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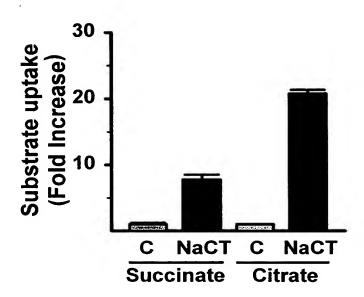


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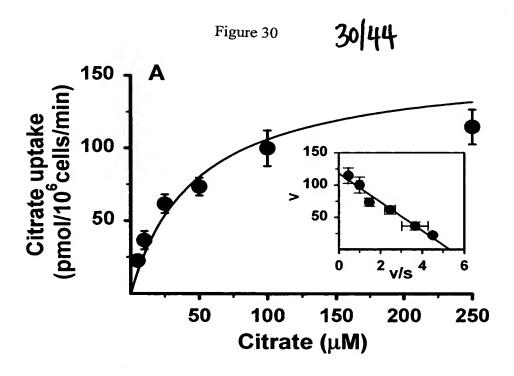


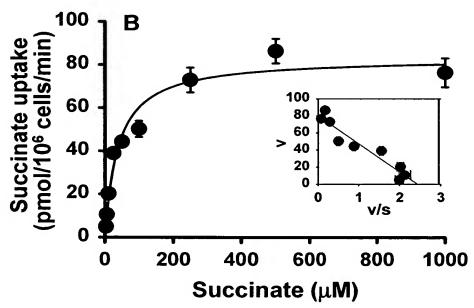
Docket: 275.00080101 Sheet 29 of 44

Figure 29



Docket: 275.00080101 Sheet 30 of 44

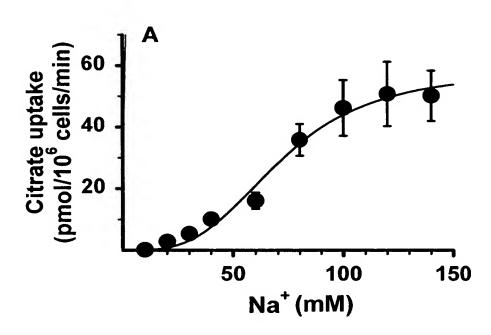


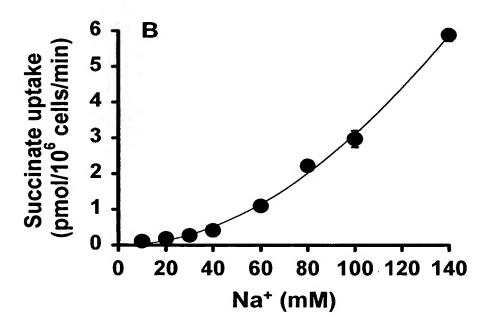


Docket: 275.00080101 Sheet 31 of 44



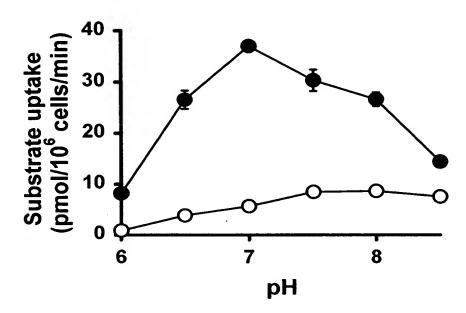
Figure 31





Docket: 275.00080101 Sheet 32 of 44

Figure 32

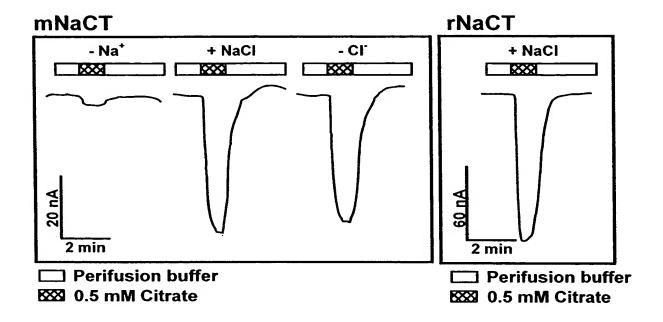


Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION Applicant(s): Ganapathy et al.

Serial No.: Unassigned Filed: Herewith
Express Mail No.: EV 201890140 US

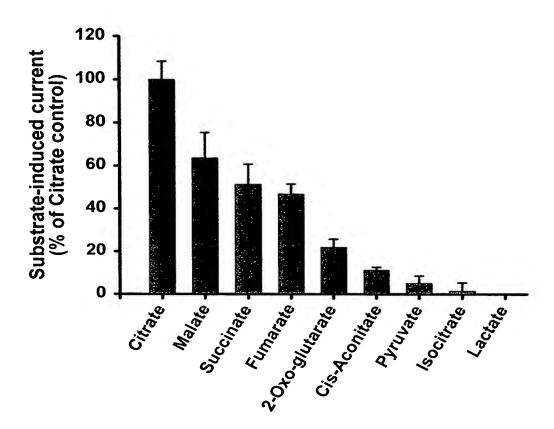
Docket: 275.00080101 Sheet 33 of 44

Figure 33

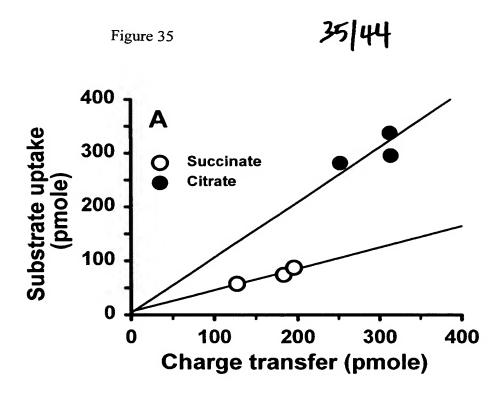


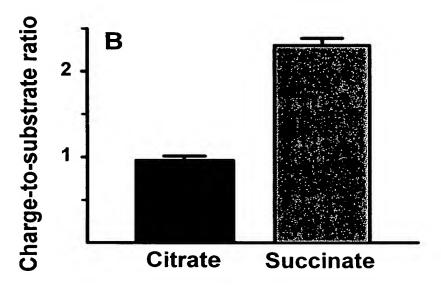
Docket: 275.00080101 Sheet 34 of 44

Figure 34



Docket: 275.00080101 Sheet 35 of 44





Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION

Applicant(s): Ganapathy et al. Serial No.: Unassigned Express Mail No.: EV 201890140 US

Filed: Herewith

Docket: 275.00080101 Sheet 36 of 44

36/44

Figure 36

Zebra Fish NaCT full length cDNA (1#) (1-2536 + 15 bp)

ORF: from 76 - 1824 (length = 1749) (SEQ IO NO 11)

AGAGATCAGCGCACAGAAGTTTTGCGCAGTTTCTCACCGTTTGGACATTTCATTGTAAAGTTATCCAAAGCCGAAA TOTALGETCACGTGCACTCAAACTAGTATGGAAGATGAAAAATACATTGATTCTTTTTTGCACTCCATTTCTTCT GCTTCCTTTGCCACTTGTCATTGGATCAAAGGAGGCTGGATGTGCATATGTTGTGGTACTGATGGCAGTTTACTGG TGTACAGAGGTGCTGCCGCTGGCTGTCACTGCTCCTCCCGCCGCTGTGCTCTTTCCCCTCTTCAGAATCATGGAGT CCCAAGACGTATGTATGCAGTACCTTAAGGACACTAACATGCTGTTTCTGGGTGGCCTGATGGTGGCCGTGGCTGT CGAACACTGGAATCTGCACAAGCGGATCGCCCTGCGGGTGCTGCTCTTGTTGGGGGGTTCGACCAGCTCTGTTAATG TTGGGCTTCATGGGTGTAACAGCTTTCCTCTCCATGTGGATCAGTAACACGGCCACAACAGCCATGATGGTGCCCA TCGTTCAGGCAGTTCTCGAGCAGCTCAACAACACAGCACAACAAGAACAAAGCTCCATACCTGAGACCGAGGAAAA GAGCACTGAGAAACAGCCTGAGAGCCCGGGTGAGGAAAAAGTGGTACTGAATGGCGACAACTTCTCAATGGAGTCA GACCCTGAAGAACATTCACGAGAAGCAGAGGAAAGGCTGAAGATGTCTAAAGGCCTGACCCTGTGCGTGTTATG CCGCCAGCATCGGCGGCACACCACACCACAGGCACTGGACCAAACCTCGTTCTTATGGGACAGATGAGCCAACT GTTCCCGGACAACCCTGACATCATTAACTTTGCGTCATGGTTTGGATTTGCCTTTCCAAACATGATCATCATGCTC ACGCTGGCCTGGCTGTGCTACAGATCGTGTTTCTGGGAATAAACTTTAAAAAGACATGGGGCTGTGGGACGGTGA AGACGGAGAAGGAGATCGCGGCCTATAATGTGATTAAAGAGGAGCACCGCAGTCTCGGCCCTATGACCTTTGGGGA GCTGAGTGTCCTTGCCCTCTTCATCCTCCTGGTGGTGCTTTGGTTCACTCGTGATCCAGGCTTCGTGGACGGCTGG GCGACACGCTTCTTCAATGCTGACAAAGAGTTTGTGACAGATGCCACGGTTGCAGTGTTTGTGGCTGCTCTCT TTGTCTTTCCCTCTAAACCACCACGATTGTGCTTCTGGAGAACAGAGGTTTCGACACAGTGCCCCAGCAAGAAAG TGGCCCGACTCCAGCTTTGCTGACATGGAAAGTGACACAGAAGAAGATGCCATGGAGTATTATACTGCTGCTGGGA GGAGGCTTTGCCCTGGCTAAGGGCAGTGAGATCTCAGGATTGTCCAAGTGGCTTGGAGATCAGATGTCTCCTCTTC AAAGCATTCCTCCATGGGCAATAGCTATTGTCATATGTTTAATGATCGCAACCTTCACTGAATGCACCAGTAATGT GGCCACAGCTACATTATTTCTGCCTATACTGGCATCTATGTCTCAGTCTATAGGTGTGAATCCTCTGTATGTTATG GTGCCCTGTACCCTCAGTGCATCTTTTGCCTTCATGCTCCCTGTGGCAACTCCTCCAAACGCCATCGTCTTCTCAT ACGGATACCTCAAAGTCTCTGACATGGCCAAGACTGGGATCGTCATGAACATCATCGGCATCCTCTCCATCACCTT AGCCATTAACAGCTGGGGCAGAGCCATCTTCAGTTTAGACACGTTCCCCAGCTGGGCAAACACTACTGATGTC GAGACACAGAGAGCCGGACTGCCCCACTCACCACTTGTGAACTTCAGATTGTTTCCAGTTCTCATGTGAACAGAGA AAACCGATTATGACCACTGTTTAGTCATTTCACACATTCATGTCTATCTTTTACAAAACCGTGGTGTTCACTACAG TCTGTGGACATTGTAACACCAATATTTGAATGATTCATAAAAGACAAAAATAGATTTTACAAATCATGATTTTTCT GTAGCATTACAGCATACTGTGAGCACTGAGCATATATTTGGACCATTGGTTGTTCGCTTTTGGCTTGTGAAAGAGC ATTTGGACGCAGAAACTCAACATCAGTCATATTGTCCCCTTGTGTCCAGACTCAGAGCCAGTGGCCGGTTTCACTG GAGATCAATACTGTACTTTGACGGTTCAAATCACTTCATGTTCATATGCGTAATTTAAAGCTGCTTTAAACACAGC TACACAAGAGCACATGCAGAAAAAGCTGAAAAGGCTGAAATTGTGTAAATATCATCTAGATTTTTAAGCACAAAAGT ATCATGTACATTTCAAAATTAATTCAAACTCTATTTTTAATGTACATTTATTAAAATTATGTATTTTGTTCATATT TACTELEGATTTTTATTTGGTTTACTC ΑΑΑΑΑΑΑΑΑΑΑΑ

Amino Acid Sequence (581 aa) (SEQ IÛ NO: 12)

MASRALKLVWKMKNTLILFCTPFLLLPLPLVIGSKEAGCAYVVVLMAVYWCTEVLPLAVTALLPAVLFPLFRIMES QDVCMQYLKDTNMLFLGGLMVAVAVEHWNLHKRIALRVLLLVGVRPALLMLGFMGVTAFLSMWISNTATTAMMVPI VQAVLEQLNNTAQQEQSSIPETEEKSTEKQPESPGEEKVVLNGDNFSMESDPEEHSREAEERLKMSKGLTLCVCYA ASIGGTATLTGTGPNLVLMGQMSQLFPDNPDIINFASWFGFAFPNMIIMLTLAWLWLQIVFLGINFKKTWGCGTVK TEKEIAAYNVIKEEHRSLGPMTFGELSVLALFILLVVLWFTRDPGFVDGWATRFFNADKEFVTDATVAVFVAALLF VFPSKPPRLCFWRTESFDTVPQQESGPTPALLTWKVTQKKMPWSIILLLGGGFALAKGSEISGLSKWLGDQMSPLQ SIPPWAIAIVICLMIATFTECTSNVATATLFLPILASMSQSIGVNPLYVMVPCTLSASFAFMLPVATPPNAIVFSY GYLKVSDMAKTGIVMNIIGILSITLAINSWGRAIFSLDTFPSWANTTDV

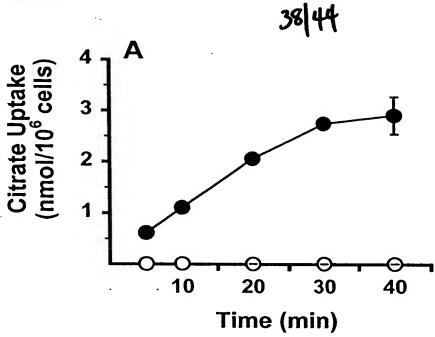
Docket: 275.00080101 Sheet 37 of 44

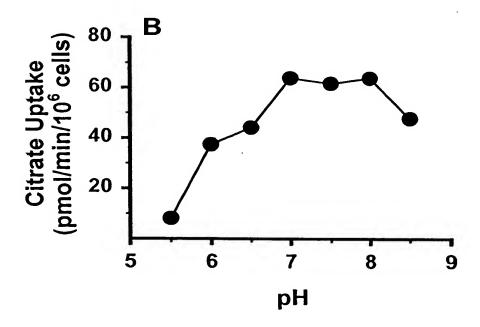
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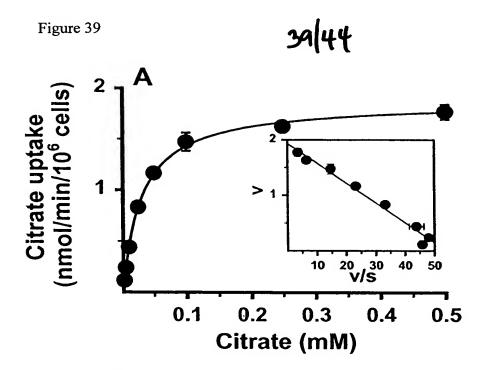
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fish mouse rat human	61 60 60	ALLF <mark>AVLFPLFRIMESOD</mark> VCMQYLKDTNMLFLGCLMVAVAVEHWNLHKRIALRVLLLVGV SLLPVLLFPLLKVLDSKQVCIQYMKDTNMLFLGSLIVAVAVERWKLHKRVALRMLLFVGT SLLPVLLFPLLKVLDSKQVCVQYMLDTNMLFLGSLIVATAVERWBLHKRIALRMLLFVGT SLMPVLLFPLFOTLDSRQVCVQYMKDTNMLFLGGLIVAVAVERWNLHKRIALRTLLWVGA
fish mouse rat human	121 120 120 120	RPALLMLGFMGVTAFLSMVISNTATTAMMVPIVQAVLBCLMNTAQQEQSSIPETBEKSTE KESRLMLGFMFVTAFLSMVISNTA <mark>A</mark> TAMMIPIVEAMLQQMIAAMTAVEASLGTLBLLDKN KESRLMLGFMFVTAFLSMVISNTATTAMMIPIVEAMLBQMVATNVAVDASORIMBLLDKN KPARLMLGFMGVTA <mark>L</mark> LSMVISNTATTAMMVPIVEAÏLQQMBATSARTBAGLELVDMG
fish mouse rat human	181 180 180 177	KOPESPGBEKVVLNGDNFSMESDFBEHSREAEERLKMSKGLTLCVCYAASIGGTATLTGT KISELPGSOVVFEDPNVCDOEDEETKNMYKAMHLCVCYŠASIGGTATLTGT KASELPGSOVVFEDPSVCKOEDEETKNMYKAMNLCVCYAASIGGTATLTGT KAKELPGSOVVFEGPTLGOOEDOERKRUCKAMTLCICYAASIGGTATLTGT
fish mouse rat human	241 231 231 228	GPNVVLLGQMQELFPDSKDVMNFASWFAFALPNMLLMLVMAWLWLLGFYMRPNLKKICIC
fish mouse rat human	291 291	G-TVKTEKELAAYNVIKBEHRSLGEMTFGELSVLALFILLVYLWFTRDPGFVDGWATR-F CGEKKRDTEKIAYKVLNEEYOKLGSLSYPECNVLFCFTLLVILWFSRDPGFMPGWLSFAX CGRKKKDTEKIASKVLYEEYRKLGPLSYAECNVLFCFGLLIILWFSRDPGFMPGWLSIAW GLESKK-NEKAALKVLQEEYRKLGPLSFAETNVLICFFLLVILWFSRDPGFMPGWLTVAW
fish mouse rat human	359 351 351 347	VEGNTVHITDATVAIFVAILLFIJPSQKPKFNFSSQTEEERFTPFYPFALLDWKVAQ LEGNTKHVTDATVAIFVAILLFIVPSQKPKFNFSRQTEEERFTPFYPFPLLNWKVTQ
fish mouse rat human	419 408 408 404	EKVPWDIVLLLGGGFAMAKGCETSGLSKWMA <mark>A</mark> CMEPLRLVKPAVITLILSCLVAMTTECT EKVPWGIVLLLGGGFAMAKGCETSGLSEWMARQMEPLSSVRPAIITLILSCIVAMTTECT
fish mouse rat human	479 468 468 464	The state of the s
fish mouse rat human	539 528 528 524	DMMKTGLIMM <mark>e</mark> Vgilsvflsvntwgramfnle <mark>n</mark> fpdwam <mark>sts</mark> vnt DMVKTGLVMNILGI <mark>a</mark> svflsvntwgravfnldkfpdwamlthint

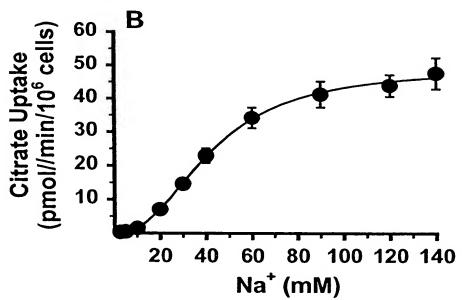
Docket: 275.00080101 Sheet 38 of 44





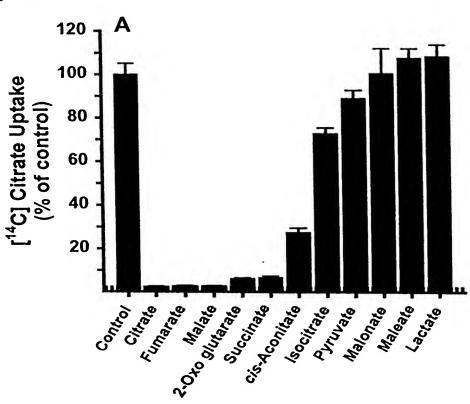


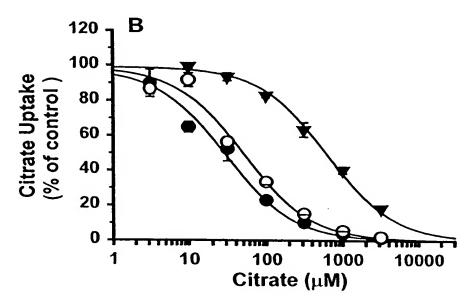




Docket: 275.00080101 Sheet 40 of 44





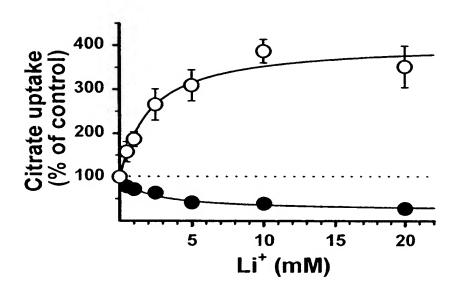


Title: NaCT AS A TARGET FOR LIFESPAN EXPANSION AND WEIGHT REDUCTION Applicant(s): Ganapathy et al.

Serial No.: Unassigned Filed: Herewith Express Mail No.: EV 201890140 US

Docket: 275.00080101 Sheet 41 of 44

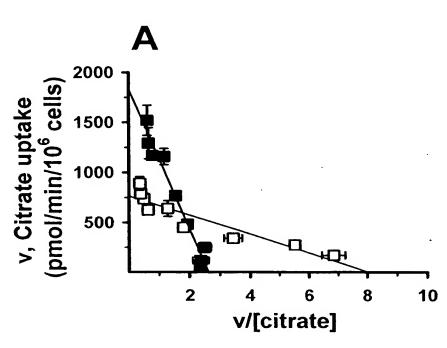
Figure 41

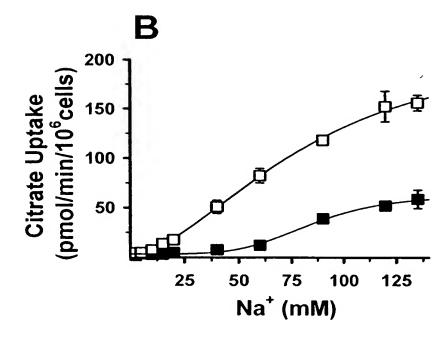


Docket: 275.00080101 Sheet 42 of 44



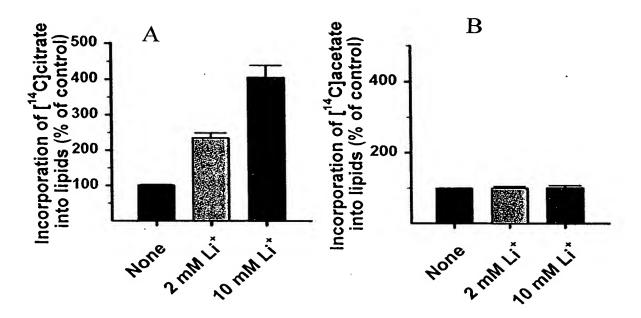






Docket: 275.00080101 Sheet 43 of 44

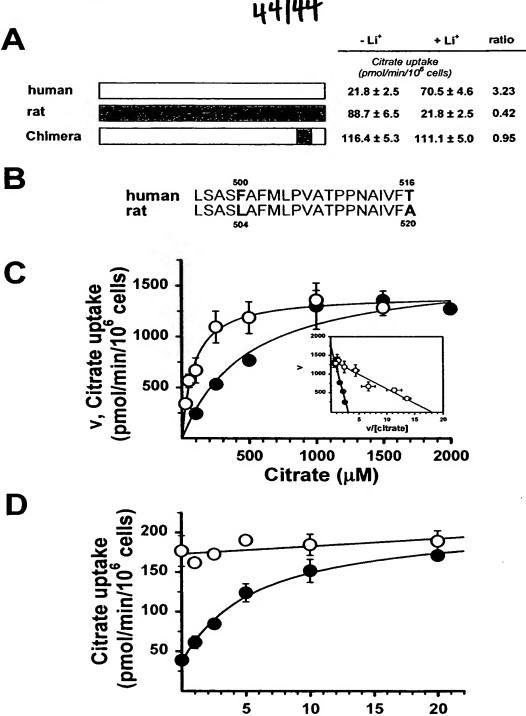
Figure 43



Docket: 275.00080101 Sheet 44 of 44



44/44



Li⁺ (mM)